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* * * * * Welcome to STN International * * * * *

NEWS	1		Web Page URLs for STN Seminar Schedule - N. America
NEWS	2	Sep 17	IMSworld Pharmaceutical Company Directory name change to PHARMASEARCH
NEWS	3	Oct 09	Korean abstracts now included in Derwent World Patents Index
NEWS	4	Oct 09	Number of Derwent World Patents Index updates increased
NEWS	5	Oct 15	Calculated properties now in the REGISTRY/ZREGISTRY File
NEWS	6	Oct 22	Over 1 million reactions added to CASREACT
NEWS	7	Oct 22	DGENE GETSIM has been improved
NEWS	8	Oct 29	AAASD no longer available
NEWS	9	Nov 19	New Search Capabilities USPATFULL and USPAT2
NEWS	10	Nov 19	TOXCENTER(SM) - new toxicology file now available on STN
NEWS	11	Nov 29	COPPERLIT now available on STN
NEWS	12	Nov 29	DWPI revisions to NTIS and US Provisional Numbers
NEWS	13	Nov 30	Files VETU and VETB to have open access
NEWS	14	Dec 10	WPINDEX/WPIDS/WPIX New and Revised Manual Codes for 2002
NEWS	15	Dec 10	DGENE BLAST Homology Search
NEWS	16	Dec 17	WELDASEARCH now available on STN
NEWS	17	Dec 17	STANDARDS now available on STN
NEWS	18	Dec 17	New fields for DPCI
NEWS	19	Dec 19	CAS Roles modified
NEWS	20	Dec 19	1907-1946 data and page images added to CA and CAPlus
NEWS	21	Jan 25	BLAST(R) searching in REGISTRY available in STN on the Web
NEWS	22	Jan 25	Searching with the P indicator for Preparations
NEWS	23	Jan 29	FSTA has been reloaded and moves to weekly updates
NEWS	24	Feb 01	DKILIT now produced by FIZ Karlsruhe and has a new update frequency
NEWS	25	Feb 19	Access via Tymnet and SprintNet Eliminated Effective 3/31/02
NEWS EXPRESS			February 1 CURRENT WINDOWS VERSION IS V6.0d, CURRENT MACINTOSH VERSION IS V6.0a(ENG) AND V6.0Ja(JP), AND CURRENT DISCOVER FILE IS DATED 05 FEBRUARY 2002
NEWS HOURS			STN Operating Hours Plus Help Desk Availability
NEWS INTER			General Internet Information
NEWS LOGIN			Welcome Banner and News Items
NEWS PHONE			Direct Dial and Telecommunication Network Access to STN
NEWS WWW			CAS World Wide Web Site (general information)

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FILE COVERS 1907 - 28 Feb 2002 VOL 136 ISS 9

FILE LAST UPDATED: 26 Feb 2002 (20020226/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

CAS roles have been modified effective December 16, 2001. Please check your SDI profiles to see if they need to be revised. For information on CAS roles, enter HELP ROLES at an arrow prompt or use the CAS Roles thesaurus (/RL field) in this file.

The P indicator for Preparations was not generated for all of the CAS Registry Numbers that were added to the CAS files between 12/27/01 and 1/23/02. As of 1/23/02, the situation has been resolved. Searches and/or SDIs in the H/Z/CA/CAPLUS files incorporating CAS Registry Numbers with the P indicator executed between 12/27/01 and 1/23/02 may be incomplete. See the NEWS message on this topic for more information.

=> s thiol?(w)chitosan

67200 THIOL?

12013 CHITOSAN

629 CHITOSANS

12031 CHITOSAN

(CHITOSAN OR CHITOSANS)

L1 3 THIOL?(W)CHITOSAN

=> d L1 1-3 ti

L1 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2002 ACS

TI Synthesis and in vitro evaluation of chitosan-thioglycolic acid conjugates

L1 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2002 ACS

TI Thiolated polymers - thiomers: development and in vitro evaluation of chitosan-thioglycolic acid conjugates

L1 ANSWER 3 OF 3 CAPLUS COPYRIGHT 2002 ACS
TI Synthesis and in vitro evaluation of chitosan-cysteine conjugates

=> d L1 1-3 ibib,abs

L1 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2001:600181 CAPLUS
TITLE: Synthesis and in vitro evaluation of
chitosan-thioglycolic acid conjugates
AUTHOR(S): Bernkop-Schnurch, Andreas; Hopf, Thorid E.
CORPORATE SOURCE: Institute of Pharmaceutical Technology and
Biopharmaceutics, Center of Pharmacy, University of
Vienna, Vienna, A-1090, Austria
SOURCE: Sci. Pharm. (2001), 69(2), 109-118
CODEN: SCPHA4; ISSN: 0036-8709
PUBLISHER: Oesterreichische Apotheker-Verlagsgesellschaft
DOCUMENT TYPE: Journal
LANGUAGE: English
AB The cationic thiomers chitosan-thioglycolic acid (TGA) shows excellent
mucoadhesive features. In order to deepen the knowledge concerning this
new excipient the optimization of its synthesis and a detailed
characterization of its properties was the objective of this study
Mediated by increasing quantities of a carbodiimide, thioglycolic acid was
covalently attached to chitosan forming amide bonds with the primary
amino
groups of the polymer Detd. with Ellman's reagent, 38 .+- . 3, 104 .+- . 2,
685 .+- . 43, and 885 .+- . 7 .mu.mol thiol groups (n=3, .+- . SD) were
bound
per g polymer at carbodiimide concns. of 50, 75, 100, and 125 mM, resp.
The immobilized thiol groups displayed a comparatively higher reactivity
to form disulfide bonds than the thiol groups in a corresponding mixt. of
chitosan and free unconjugated TGA. In an aq. 0.5% (m/v) chitosan-TGA
gel
59 .+- . 5% of the thiol groups formed disulfide bonds within 6 h at pH
6.0, whereas merely 5 .+- . 3% were oxidized in the corresponding phys.
mixt. of chitosan and TGA. Diffusion studies showed that the modified
polymer was capable of binding cysteine and cysteine Me ester. The
result
supports the theory that the improved mucoadhesive properties of
thiolated chitosan are based on the formation of
disulfide bonds with cysteine moieties of mucus glycoproteins. Because
of
its availability via an efficient synthetic pathway and its mucoadhesive
properties based on the capability to bind cysteine subunits,
chitosan-TGA
seems to be a promising new excipient for various drug delivery systems.
REFERENCE COUNT: 17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR
THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE
FORMAT

L1 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2001:520176 CAPLUS
TITLE: Thiolated polymers - thiomers: development and in
vitro evaluation of chitosan-thioglycolic acid
conjugates
AUTHOR(S): Kast, C. E.; Bernkop-Schnurch, A.
CORPORATE SOURCE: Center of Pharmacy, Institute of Pharmaceutical

Vienna, Technology and Biopharmaceutics, University of

Vienna, A-1090, Austria
SOURCE: Biomaterials (2001), 22(17), 2345-2352
CODEN: BIMADU; ISSN: 0142-9612

PUBLISHER: Elsevier Science Ltd.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The aim of this study was to improve mucoadhesive properties of chitosan by the covalent attachment of thiol moieties to this cationic polymer. Mediated by a carbodiimide, thioglycolic acid (TGA) was covalently attached to chitosan. This was achieved by the formation of amide bonds between the primary amino groups of the polymer and the carboxylic acid group of TGA. Dependent on the pH-value and the wt. ratio of polymer to TGA during the coupling reaction the resulting thiolated polymers, the so-called thiomers, displayed 6.58, 9.88, 27.44, and 38.23 .mu.mole thiol groups per g polymer. Tensile studies carried out with these

chitosan-TGA

conjugates on freshly excised porcine intestinal mucosa demonstrated a 6.3-, 8.6-, 8.9-, and 10.3-fold increase in the total work of adhesion (TWA) compared to the unmodified polymer, resp. In contrast, the combination of chitosan and free unconjugated TGA showed almost no mucoadhesion. These data were in good correlation with further results obtained by another mucoadhesion test demonstrating a prolonged residence time of **thiolated chitosan** on porcine mucosa. The swelling behavior of all conjugates was thereby exactly in the same range as for an unmodified polymer pretreated in the same way. Furthermore, it could be shown that chitosan-TGA conjugates are still biodegradable by

the

glycosidase lysozyme. According to these results, chitosan-TGA

conjugates

represent a promising tool for the development of mucoadhesive drug delivery systems.

REFERENCE COUNT: 24 THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE
FORMAT

L1 ANSWER 3 OF 3 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2000:31626 CAPLUS

DOCUMENT NUMBER: 132:98016

TITLE: Synthesis and in vitro evaluation of
chitosan-cysteine

conjugates

AUTHOR(S): Bernkop-Schnurch, Andreas; Brandt, Ursula-Maria;
Clausen, Andreas E.

CORPORATE SOURCE: Institut Pharmazeutische Technologie,
Pharmazie-Zentrum, Univ. Wien, Vienna, A-1090,

Austria

SOURCE: Sci. Pharm. (1999), 67(4), 197-208

CODEN: SCPHA4; ISSN: 0036-8709

PUBLISHER: Oesterreichische Apotheker-Verlagsgesellschaft

DOCUMENT TYPE: Journal

LANGUAGE: German

AB Mediated by a water-sol. carbodiimide cysteine was covalently attached to chitosan. According to the amt. of carbodiimide during the coupling reaction, 0.25, 0.7, and 1.2% of Cys were thereby bound to the polymer. Whereas the mucoadhesive properties of chitosan could not be improved due to this modification, the stability of matrix tablets based on **thiolated chitosan** might be strongly improved because of

the formation of inter- and/or intramol. disulfide bonds within these polymers. This oxidative process can be accelerated at higher temps. and by lowering the proton concn. on the polymer. Permeation studies carried out by chambers with freshly excised intestinal mucosa from guinea pigs demonstrated furthermore an improved permeation enhancing effect of chitosan due to the covalent attachment of Cys on it.

REFERENCE COUNT: 11 THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS
RECORD. ALL CITATIONS AVAILABLE IN THE RE
FORMAT

=> s thiol?(w)pectin
67200 THIOL?
17317 PECTIN
4168 PECTINS
18878 PECTIN
(PECTIN OR PECTINS)
L2 0 THIOL? (W) PECTIN

=> s thiol?(w)hyaluronic
67200 THIOL?
10110 HYALURONIC
1 HYALURONICS
10110 HYALURONIC
(HYALURONIC OR HYALURONICS)
L3 0 THIOL? (W) HYALURONIC

=> s thiol?(w)carboxymethylcellulose
67200 THIOL?
5091 CARBOXYMETHYLCELLULOSE
48 CARBOXYMETHYLCELLULOSES
5111 CARBOXYMETHYLCELLULOSE
(CARBOXYMETHYLCELLULOSE OR CARBOXYMETHYLCELLULOSES)
L4 1 THIOL? (W) CARBOXYMETHYLCELLULOSE

=>

=> d L4 ti

L4 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2002 ACS
TI **Thiolated carboxymethylcellulose:** in vitro evaluation
of its permeation enhancing effect on peptide drugs

=> d l4 ibib

L4 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2001:24250 CAPLUS
DOCUMENT NUMBER: 135:308715
TITLE: **Thiolated carboxymethylcellulose:**
effect in vitro evaluation of its permeation enhancing
on peptide drugs
AUTHOR(S): Clausen, A. E.; Bernkop-Schnurch, A.
CORPORATE SOURCE: Institute of Pharmaceutical Technology and
Biopharmaceutics, Centre of Pharmacy, University of
Vienna, Austria
SOURCE: Eur. J. Pharm. Biopharm. (2001), 51(1), 25-32
CODEN: EJPBEL; ISSN: 0939-6411

PUBLISHER: Elsevier Science Ireland Ltd.
DOCUMENT TYPE: Journal
LANGUAGE: English
REFERENCE COUNT: 28 THERE ARE 28 CITED REFERENCES AVAILABLE FOR
THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE
FORMAT

=> s thiol?(w)hydroxypropylcellulose
67200 THIOL?
1581 HYDROXYPROPYLCELLULOSE
4 HYDROXYPROPYLCELLULOSES
1582 HYDROXYPROPYLCELLULOSE
(HYDROXYPROPYLCELLULOSE OR HYDROXYPROPYLCELLULOSES)
L5 0 THIOL? (W)HYDROXYPROPYLCELLULOSE

=> s thiol? and hydroxypropylcellulose
67200 THIOL?
1581 HYDROXYPROPYLCELLULOSE
4 HYDROXYPROPYLCELLULOSES
1582 HYDROXYPROPYLCELLULOSE
(HYDROXYPROPYLCELLULOSE OR HYDROXYPROPYLCELLULOSES)
L6 3 THIOL? AND HYDROXYPROPYLCELLULOSE

=> d L6 1-3 ti,kwic

L6 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2002 ACS
TI Curly hair-straightening composition comprising keratin-reducing
substances and alcohols
AB . . . sufficiently straighten natural curly or frizzy hair without
damaging the hair. Thus, a 1-pack product contained NaHSO₃ 2.0,
2-methyl-2,4-pentanediol 30.0, **hydroxypropylcellulose** 1.5, water
64.0 wt.%, and monoethanolamine to adjust the pH to 9.5.
ST hair straightening compn **thiol** alc

L6 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2002 ACS
TI Tetrahydro-5-oxo-2-(3-oxo-2-isoxazolidinyl)-2-furancarboxylates
AB . . . bacteria, e.g., Staphylococcus aureus with min. inhibitory
concn.
of 6.25 .mu.g/mL. Tablets contg. II 300, corn starch 50, lactose 28,
hydroxypropylcellulose L 20 and Mg stearate 2 mg were prepd.
IT 75-08-1, Ethanethiol 108-98-5P, Thiophenol, preparation
RL: RCT (Reactant)
(**thiolation** by, of bromooxoglutaric acid)
IT 89469-94-3
RL: RCT (Reactant)
(**thiolation** of, by thiophenol)

L6 ANSWER 3 OF 3 CAPLUS COPYRIGHT 2002 ACS
TI **Thiol** derivatives of cellulose as supports for the
immobilization of non-**thiol** enzymes
TI **Thiol** derivatives of cellulose as supports for the
immobilization of non-**thiol** enzymes
AB . . . were introduced into cellulose in 2 ways. Chlorodeoxycellulose
gave with Na₂S₂O₃ a thiosulfate deriv., which was reduced to
mercaptocellulose. Similarly, 3-chloro-2-**hydroxypropylcellulose**
was converted to epoxide, which on reaction with Na₂S₂O₂ and redn. gave
3-mercapto-2-**hydroxypropylcellulose**. Acetylcholinesterase (EC
3.1.1.7) from bovine erythrocytes and from elec. eel,

butyrylcholinesterase (EC 3.1.1.8), and trypsin (EC 3.4.21.4) were immobilized on. . .

ST enzyme immobilization cellulose **thiol** deriv;
 acetylcholinesterase immobilization cellulose **thiol** deriv;
 cholinesterase immobilization cellulose **thiol** deriv; trypsin
 immobilization cellulose **thiol** deriv

IT Enzymes
 RL: PROC (Process)
 (immobilization of, on cellulose **thiol** derivs.)

IT 9000-81-1 9001-08-5 9002-07-7
 RL: PROC (Process)
 (immobilization of, on cellulose **thiol** derivs.)

IT 9001-08-5DP, mercaptobutylamidino deriv. 9002-07-7DP,
 isothiocyanatopropyl deriv.
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (prepn. and immobilization on cellulose **thiol** derivs.)

IT 9004-34-6DP, **thiol** derivs. 37324-27-9P
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (prepn. and use in enzyme immobilization)

=> d L6 3 ibib

L6 ANSWER 3 OF 3 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 1981:564692 CAPLUS
 DOCUMENT NUMBER: 95:164692
 TITLE: **Thiol** derivatives of cellulose as supports
 for the immobilization of non-**thiol** enzymes
 AUTHOR(S): Gemeiner, Peter; Zemek, Jiri
 CORPORATE SOURCE: Inst. Chem., Slovak Acad. Sci., Bratislava, 809 33,
 Czech.
 SOURCE: Collect. Czech. Chem. Commun. (1981), 46(7), 1693-700
 CODEN: CCCCAK; ISSN: 0366-547X
 DOCUMENT TYPE: Journal
 LANGUAGE: English

=> s thiol? and pectin
 67200 THIOL?
 17317 PECTIN
 4168 PECTINS
 18878 PECTIN
 (PECTIN OR PECTINS)

L7 18 THIOL? AND PECTIN

=> d L18 1-18 ti
 L18 NOT FOUND
 The L-number entered has not been defined in this session, or it
 has been deleted. To see the L-numbers currently defined in this
 session, enter DISPLAY HISTORY at an arrow prompt (=>).

=> d L7 1-18 ti

L7 ANSWER 1 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI Adsorbent preparation, dry process for crosslinked polysaccharides, and
 recovery and separation of arsenic ions using the same

L7 ANSWER 2 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI Oral pharmaceutical preparation embedded in an oily matrix and coated by
 enteric coating polymers

- L7 ANSWER 3 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI Kiwi protein inhibitor of **pectin** methylesterase. Amino-acid sequence and structural importance of two disulfide bridges
- L7 ANSWER 4 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI Purification and some properties of protease from *Actinidia eriantha* Benth
- L7 ANSWER 5 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI Keratinocyte growth factor-2 formulations for promotion of wound healing
- L7 ANSWER 6 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI Status of the **thiol**-dependent cytoprotectant systems under conditions of lead intoxication and high-**pectin** diet
- L7 ANSWER 7 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI Effect of mercurials on the activity of **pectin** methylesterase from egg plant (*Solanum melongena* L.)
- L7 ANSWER 8 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI Pharmaceutical compositions containing biologically active agents contained within a polymeric shell
- L7 ANSWER 9 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI Stabilized suspension of magnetic particles and its preparation and use in NMR diagnosis
- L7 ANSWER 10 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI Characterization of long-term extension of isolated cell walls from growing cucumber hypocotyls
- L7 ANSWER 11 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI Purification and properties of actinidin from *Actinidia chinensis*
- L7 ANSWER 12 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI An alkaline extracellular protease produced by *Cladosporium cucumerinum* and its possible importance in the development of scab disease of cucumber seedlings
- L7 ANSWER 13 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI Deodorization by aerobes
- L7 ANSWER 14 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI **Pectin** esterase(s) from sour oranges (*Citrus aurantium* Linn)
- L7 ANSWER 15 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI Rust remover containing a mercapto compound
- L7 ANSWER 16 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI Colorimetric determination of volatile sulfur compounds in foods. II. Reaction with bis(p-nitrophenyl) disulfide
- L7 ANSWER 17 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI Origin of methanol and dimethyl sulfide from cooked foods
- L7 ANSWER 18 OF 18 CAPLUS COPYRIGHT 2002 ACS
 TI Reductones derived from 3,4-dihydroxy-2,5-dicarboxylic esters of furan,

thiophene, N-phenylpyrrole, and selenophene

=> s thiol?(p)hyaluronic

67200 THIOL?

10110 HYALURONIC

1 HYALURONICS

10110 HYALURONIC

(HYALURONIC OR HYALURONICS)

L8 21 THIOL?(P)HYALURONIC

=> d L8 1-21 ti,kwic

L8 ANSWER 1 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI Gold plating and biofunctionalization of ferromagnetic magnetic tweezers:
Application for local studies of soft surface-grafted polymer films
AB . . . resulting gold layer is a versatile platform for further
biofunctionalization using a wide variety of std. coupling protocols
based

on **thiol** chem. Several methods, such as electron microscopy,
elemental anal., x-ray powder diffraction, and XPS, have been used for
quant. and. . . of the coatings. The magnetic tweezers are used for
local quant. characterization of the elasticity of soft surface-grafted
films of **hyaluronic** acid. A method for the calibration of the
magnetization of each bead chosen for the measurement is introduced which
is. . .

L8 ANSWER 2 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI Hyaluronic acid-protein conjugates, pharmaceutical compositions and
related methods

AB . . . present invention broadly relates to the field of protein
modification and, more specifically, the attachment of low mol. wt.,
derivatized **hyaluronic** acid polymer to proteins including leptin
and analogs thereof (the term "protein" as used herein is synonymous with
"polypeptide" or "peptide" unless otherwise indicated). The
hyaluronic acid-protein conjugates of the present invention
exhibit longer sustained blood levels than formulations contg. protein
alone, thus providing an important advantage in the therapeutic setting.
Conjugates of low mol. wt. sodium hyaluronate with free-**thiol**
osteroprotegrin was prepd.

L8 ANSWER 3 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI Synthesis of .beta.-D-glucopyranosyl(1.fwdarw.3)-1-**thiol**
-.beta.-glucosamine disaccharide derivative as building block for the
synthesis of **hyaluronic** acid

TI Synthesis of .beta.-D-glucopyranosyl(1.fwdarw.3)-1-**thiol**
-.beta.-glucosamine disaccharide derivative as building block for the
synthesis of **hyaluronic** acid

ST glucopyranosyl **thiolglucosamine** prepn **hyaluronic** acid
building block

IT Glycosylation

(synthesis of .beta.-D-glucopyranosyl(1.fwdarw.3)-1-**thiol**
-.beta.-glucosamine, a building block for the synthesis of
hyaluronic acid)

IT 9004-61-9P, **Hyaluronic** acid

RL: PNU (Preparation, unclassified); PREP (Preparation)

(synthesis of .beta.-D-glucopyranosyl(1.fwdarw.3)-1-**thiol**
-.beta.-glucosamine, a building block for the synthesis of
hyaluronic acid)

IT 66-84-2, D-Glucosamine hydrochloride 76-03-9, Trichloroacetic acid,

reactions 83-87-4, D-Glucose pentaacetate 98-88-4, Benzoyl chloride 106-45-6, p-Thiocresol 116-11-0 123-76-2, Levulinic acid 1125-88-8, Benzaldehyde dimethyl acetal 17341-93-4

RL: RCT (Reactant)

(synthesis of .beta.-D-glucopyranosyl(1.fwdarw.3)-1-**thiol**-.beta.-glucosamine, a building block for the synthesis of **hyaluronic acid**)

IT 1152-39-2P 28244-94-2P 97562-23-7P 122210-01-9P 219518-19-1P
220645-20-5P 323195-38-6P 323195-39-7P 323195-40-0P 323195-41-1P
323195-42-2P 323195-43-3P 323195-44-4P 323195-45-5P 323195-46-6P
323195-47-7P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation)
(synthesis of .beta.-D-glucopyranosyl(1.fwdarw.3)-1-**thiol**-.beta.-glucosamine, a building block for the synthesis of **hyaluronic acid**)

IT 323195-37-5P

RL: SPN (Synthetic preparation); PREP (Preparation)
(synthesis of .beta.-D-glucopyranosyl(1.fwdarw.3)-1-**thiol**-.beta.-glucosamine, a building block for the synthesis of **hyaluronic acid**)

L8 ANSWER 4 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI Oxidative damage of eye tissues and protection by thioctic acid

AB . . . in the mol. wt. compn. of bovine-lens homogenates mediated by illuminated riboflavin. Another indicator detected was the amt. of free **thiol** groups. Only the reduced form of thioctic acid (dihydrothioctic acid) protected from photo-oxidative changes, comparable to the synthetic dithiol dithiothreitol. . . by the oxidative fragmentation of .alpha.-keto-.gamma.-methiol-butyric acid (KMB) yielding ethylene. Both, thioctic acid and its reduced form, decreased ethylene formation. **Hyaluronic acid** is an important structural glycan in the vitreous. Oxidative degrdn. of **hyaluronic acid** by hypochlorous acid or Fenton-type oxidants is also diminished by thioctic acid, measured by mol.-wt. anal. of **hyaluronic acid**.

L8 ANSWER 5 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI Thiol-containing biomaterials for medical and pharmaceutical use

IT 9004-34-6, Cellulose, biological studies 9004-54-0, Dextran, biological studies 9004-61-9, **Hyaluronic acid** 9005-25-8, Starch, biological studies 9007-28-7, Chondroitin sulfate 9012-76-4, Chitosan 26780-50-7, Poly(glycolide-lactide)

RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(**thiol**-contg. biomaterials for medical and pharmaceutical use)

L8 ANSWER 6 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI Synthetic polysulfated hyaluronic acid is a potent inhibitor for tumor necrosis factor production

AB . . . blocks for synthesizing nontoxic drugs for suppression of tumor necrosis factor (TNF) prodn. by inflammatory cells, we have chem. modified

hyaluronic acid (HA) and tested its effects in blocking TNF-.alpha. and TNF-.beta. prodn. in vitro. HA was chosen mainly for its.

. . . decreasing the extent of polysulfation, the inhibitory effect of HAS

on TNF-.alpha. prodn. was diminished. Other chem. modifications, including deacetylation, **thiolation**, or redn. of the carboxylic groups, could not increase the efficacy of HA in suppression of

TNF-.alpha. prodn. Naturally polysulfated. . .

L8 ANSWER 7 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI Inhibitory actions of **thiol** compounds in HOCl induced degradation of **hyaluronic acid**

TI Inhibitory actions of **thiol** compounds in HOCl induced degradation of **hyaluronic acid**

AB Effects of **thiol** compds. N-(N-L-.gamma.-glutamyl-L-cysteinyl)glycine, N-(2-mercaptopropionyl)glycine, and cysteine on the degrdn. of **hyaluronic acid** were investigated. Scavenging actions of **thiol** compds. on HOCl were examd.

L8 ANSWER 8 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI Endothelial cell stimulation of smooth muscle glycosaminoglycan synthesis can be accounted for by transforming growth factor beta activity

AB . . . Here it is shown that the factor responsible is transforming growth factor beta (TGF-.beta.), as assessed by (1) proteinase and **thiol** sensitivity, (2) heat and acid enhancement of ECCM activity, and (3) neutralization of ECCM activity by anti-TGF-.beta. Ig. Anti-TGF-.beta. neutralization. . . showed that ECCM from EC of

varying densities stimulated individual GAG to varying degrees. ECCM from low-d. EC preferentially stimulated **hyaluronic acid** (HA), whereas ECCM from intermediate- and high-d. cultures stimulated increasing amts. of sulfated GAG. Exposure of SMC to varying. . .

L8 ANSWER 9 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI The mechanism of chondrocyte hydrogen peroxide damage. Depletion in intracellular ATP due to suppression of glycolysis caused by oxidation of glyceraldehyde-3-phosphate dehydrogenase

AB . . . be due to the oxidative inactivation of glyceraldehyde-3-phosphate dehydrogenase (G-3-PDH). Apparently, intrachondrocyte oxidant damage occurs through oxidn. of the sensitive **thiol** (-SH) residue at the active center of G-3-PDH, with subsequent redn. in the

rate of glycolytic ATP synthesis and the intracellular concn. of ATP which is required for DNA, protein, proteoglycan, and **hyaluronic acid** synthesis.

L8 ANSWER 10 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI **Hyaluronic acid** degrading reactions under hyperthermic conditions and effects of **thiols**

TI **Hyaluronic acid** degrading reactions under hyperthermic conditions and effects of **thiols**

AB **Thiols** promoted the formation of highly oxidizing **hyaluronic acid** (HA)-degrading species in the presence of chelated (EDTA) Fe3+, Fe2+, and Cu2+ compds. Hyperthermia (40-44.degree.) led to an addnl. increase in the prodn. of HA-degrading species by the interaction of **thiols** with EDTA-complexed transition metals.

The most efficient **thiols** were cysteamine and cysteine; GSH, dithiothreitol, and N-acetylcysteine promoted the generation of the

highly reactive radicals to a minor extent.

IT Transition metals, compounds

RL: BIOL (Biological study)

(EDTA complexes, oxygen radical formation in presence of, hyperthermia and **thiols** effects on, **hyaluronic acid** degrdn. in relation to)

IT Fever and Hyperthermia

Thiols, biological studies

- RL: BIOL (Biological study)
(oxygen radical formation in presence of transition metal-EDTA complexes response to, **hyaluronic acid** degrdn. in relation to)
- IT 7782-44-7D, radicals
RL: FORM (Formation, nonpreparative)
(formation of, **thiols** promotion of, hyperthermia effect on, **hyaluronic acid** degrdn. in relation to)
- IT 60-00-4D, transition metal complexes 7439-89-6D, EDTA complexes
7440-50-8D, EDTA complexes 7447-39-4D, EDTA complexes 7705-08-0D,
EDTA complexes 7720-78-7D, EDTA complexes
RL: BIOL (Biological study)
(oxygen radical formation in presence of, hyperthermia and **thiols** effects on, **hyaluronic acid** degrdn. in relation to)
- L8 ANSWER 11 OF 21 CAPLUS COPYRIGHT 2002 ACS
TI Copper dependent control of the enzymic and phagocyte induced degradation of hyaluronic acid, synovial fluid and cytochrome c
AB Studies on the oxidn. of Cu-thionein by xanthine oxidase are described in detail. Cu²⁺ was released by Cu(I)-**thiolate** (chromophore of Cu-thionein) degrdn. Complete but reversible inhibition of xanthine oxidase occurred in the presence of 5 .mu.M Cu; diminished xanthine oxidase inhibition occurred with the Cu(I)-**thiolate** chromophore of yeast Cu-thionein. The xanthine oxidase-dependent oxidn. of xanthine or hypoxanthine degraded **hyaluronic acid** (a component of synovial fluid), and Cu-thionein and CuSO₄ inhibited the degrdn. Effects on bovine synovial fluid also are. . . c by phagocyte-generated hypochlorite was a measure of polymorphonuclear leukocyte (PMN) activity. Activated PMNs produce excited O species which depolymerize **hyaluronic acid** but no degrdn. was obsd. on the presence of ceruloplasmin and CuSO₄; some degrdn. was obsd. with Cu-thionein. Relations. . .
- L8 ANSWER 12 OF 21 CAPLUS COPYRIGHT 2002 ACS
TI Radiation induced depolymerization of hyaluronic acid (HA) in aqueous solutions at pH 7.4
AB Radiolytic depolymn. of **hyaluronic acid** (HA, a heteropolysaccharide) in aq. solns. under a variety of conditions demonstrates that the damaging effect of radiolytic radical. . . order OH.cntdot.>e-aq>0-2. Cysteine, penicillamine, and dithiothreitol protected against primary radiolytic species. The enzyme superoxide dismutase (SOD) and the above 3 **thiols** do not protect against the radiolytic species generated by the .lambda.-irradn. of aerated Na formate solns. The results also indicate that the reaction between CO-2 anion and **hyaluronic acid** is faster than the reaction between O-2 and **hyaluronic acid** and that CO-2 anions are not scavenged by superoxide dismutase. The results further suggest that tert-BuOH radicals interact with **hyaluronic acid** and reduce the viscosity of HA solns. Preliminary pulse radiolysis expts. do demonstrate a reaction between CO-2 radical and **hyaluronic acid**.
- L8 ANSWER 13 OF 21 CAPLUS COPYRIGHT 2002 ACS
TI Connective tissue activation. XXVIII. A connective tissue activating peptide from human urine
AB . . . be different from EGF and IgG. This urinary connective tissue-activating factor (CTAP-U) [92307-84-1] is nondialyzable, labile to protease, stable to **thiols**, heat, and acid, and has an acidic isoelec. point. Purified prepns. of CTAP-U have biol. activities that

cause human connective tissue cells to synthesize incremental amts. of ¹⁴C-labeled **hyaluronic acid** [9004-61-9], [³⁵S]proteoglycans, and [³H]DNA in vitro. The cell spectrum responsive to this substance includes human synovial cells, human chondrocytes, . . .

L8 ANSWER 14 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI Cartilage-degrading neutral proteinase secreted by Yoshida sarcoma cells. Purification and properties

AB . . . help of a new assay system for measurement of proteoglycan core protein degradn., which utilizes aminopropyl glass beads derivatized with **hyaluronic acid**. This enzyme, with a neutral pH optimum and apparent mol. wt. of .apprx.30,000, was secreted into culture medium in. . . form. It was resistant to cartilage-derived inhibitors and to .alpha.2-macroglobulin as well as to synthetic and natural inhibitors of serine, **thiol**, and carboxyl proteinases. It was inhibited by 1,10-phenanthroline and **thiols** at relatively high concns., and therefore is probably a metalloproteinase. The enzyme degraded type V collagen, types I and II. . . .

L8 ANSWER 15 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI Primary selection and study of the properties of chemical agents for protection from radiation injury

AB . . . presented for a primary selection and study of radioprotectors, and applied in studying the protective effects on erythrocytes of aminoalkyl **thiols**, aminoalkyl disulfides, aminoalkylisothiuronium compds., Bunte salts, and thiazolidines. A radiomimetic effect of lipoidal toxic compds. produced during irradiation is significantly reduced when phospholipids from animal tissues are added. Polysaccharides except **hyaluronic acid** exhibited a similar action in the erythrocyte radiomimetic model to that of phospholipids.

L8 ANSWER 16 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI Inhibition of steroid .DELTA.4-reductase by polysulfated polysaccharides and para-chloromercuribenzoic acid

AB Four heparinoids and 2 naturally occurring, high mol. wt. polysaccharides with varying contents, including heparin, chondroitin sulfate, and **hyaluronic acid**, were assayed for inhibitory effects on rat liver steroid .DELTA.4-reductase activity. All active compds. had a 15-18% S content, . . . of cortisone reductase of only 60%, while p-chloromercuribenzoate (PCMB) (5 .times. 10⁻⁴M) completely inhibited the enzyme, indicating the importance of **thiol** groups to enzymic activity. PCMB and heparin were additive in a manner which gave greater inhibition than could be demonstrated. . . .

L8 ANSWER 17 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI Depolymerization of hyaluronic acid by autoxidants and radiations

AB The effects of x-rays, .beta.-rays (32P internally), and autoxidants were compared for the aerobic depolymerization of **hyaluronic acid**. A 0.02mM soln. of ascorbic acid reduced the viscosity of **hyaluronic acid** solns. at about the same rate as 2400 rads of x-rays or .beta.-rays. The yields (G values) were low, about 0.015. Visible light also caused the degradation of **hyaluronic acid** in the presence of riboflavine as a sensitizer. A survey of compds. with possible autoxidant

properties showed that the following structures were assocd. with such activity: enediols (reduced or oxidized), quinones and hydroquinones,

many

thiols (not disulfides), m-nitrophenol, and a few metallic ions, esp. Cu⁺, Fe⁺⁺, and Sn⁺⁺. Antioxidant activity of these compds. was tested extensively for the system **hyaluronic acid**-ascorbic acid,

and some compds. were studied in the radiation systems. The effects in both systems were similar, but were. . .

L8 ANSWER 18 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI Inactivation of solutions of transforming deoxyribonucleic acid by .gamma.-rays in the presence of protective substances

AB . . . had a protective effect. The rate of I inactivation was unaffected by irradiation rates of 2.4, 8.3, and 33 kr./hr.; **hyaluronic** acid from human umbilical cord, RNA, or yeast ext. used at 1 mg./cc. had a protective effect. The protective effect. . . the original, were 1, 8, 10, 17, 57, 170, 76, 330, 719, 92, and 560 kr./hr. for control; 1 mg./cc. **hyaluronic** acid, yeast RNA, or yeast ext. 0.25 and 1.0 mg./cc. thiourea; 0.25, 1.0, and 2.5 mg./cc. cysteine; and 0.25 and 1.0 mg./cc. mercaptoethanol, resp. The protective effect of **thiols** is considered specific and different from that of the yeast ext.

L8 ANSWER 19 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI Depolymerization of hyaluronic acid by the oxidative-reductive depolymerization (ORD) reaction

AB cf. CA 54, 4692d, 24922g. **Hyaluronic** acid (I) was depolymerized in vitro by various biol. occurring reducing agents, such as ferrous ions, ascorbic acid, hydroquinones, and **thiol** compds. Depolymerization was followed by the fall in intrinsic viscosity of 40 mg. % solns. of I in 0.2M phosphate. . .

L8 ANSWER 20 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI State of polysaccharides (hyaluronic acid) in rheumatism

AB . . . the specific enzyme hyaluronidase is increased. An accumulation of nonspecific substances takes place in rheumatic patients. They cause destruction of **hyaluronic** acid and non-specific substances are excreted with the urine. These substances are found also in the urine of normal persons but in a much smaller amt. These nonspecific substances include ascorbic acid, diazo compds. and azoproteins, lecithin, **thiolactic** acid, some P compds., etc. Following treatment with salicylates the activity of hyaluronidase is depressed and the amt. of nonspecific. . .

L8 ANSWER 21 OF 21 CAPLUS COPYRIGHT 2002 ACS

TI Diffusing factors. The hyaluronidase activity of testicular extracts, bacterial culture filtrates and other agents that increase tissue permeability

AB The assocn. between hyaluronidase activity (ability to hydrolyze **hyaluronic** acid, a mucopolysaccharide present in certain mucoproteins) and diffusing factors from various sources was described. The reduction of viscosity and. . . oxidation products also possessed both these properties. Reducing substances were not formed in either case. Other reducing substances such as **thiolacetic** acid, H₂S, hydroquinone, pyrogallol, Na₂SO₃ and metol also decreased the viscosity of mucoprotein and had diffusing activity when injected. The. . .

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L8 ANSWER 5 OF 21 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1995:774579 CAPLUS

DOCUMENT NUMBER: 123:208920

TITLE: Thiol-containing biomaterials for medical and pharmaceutical use
 INVENTOR(S): Constancis, Alain; Soula, Gerard
 PATENT ASSIGNEE(S): Flamel Technologies, Fr.
 SOURCE: Fr. Demande, 28 pp.
 CODEN: FRXXBL
 DOCUMENT TYPE: Patent
 LANGUAGE: French
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
FR 2707992	A1	19950127	FR 1993-9198	19930721
FR 2707992	B1	19951013		
WO 9503272	A1	19950202	WO 1994-FR914	19940721
W: JP, US				
RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
EP 710226	A1	19960508	EP 1994-922288	19940721
EP 710226	B1	19981014		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE				
JP 09503490	T2	19970408	JP 1994-504980	19940721
AT 172191	E	19981015	AT 1994-922288	19940721
US 5646239	A	19970708	US 1996-578539	19960306
PRIORITY APPLN. INFO.:			FR 1993-9198	19930721
			WO 1994-FR914	19940721
OTHER SOURCE(S):			MARPAT 123:208920	

L8 ANSWER 6 OF 21 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 1994:570103 CAPLUS
 DOCUMENT NUMBER: 121:170103
 TITLE: Synthetic polysulfated hyaluronic acid is a potent inhibitor for tumor necrosis factor production
 AUTHOR(S): Chang, Nan Shan; Intrieri, Catherine; Mattison, Jeffery; Armand, Gerard
 CORPORATE SOURCE: Lab. Mol. Immunol., Guthrie Res. Inst., Sayre, PA, USA
 SOURCE: J. Leukocyte Biol. (1994), 55(6), 778-84
 CODEN: JLBIE7; ISSN: 0741-5400
 DOCUMENT TYPE: Journal
 LANGUAGE: English

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COST IN U.S. DOLLARS

SINCE FILE	TOTAL
ENTRY	SESSION
92.21	92.36

FULL ESTIMATED COST

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE	TOTAL
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